

LISTING OF CLAIMS

The listing of claims will replace all prior versions of claims in the application.

1. (Currently amended) A nucleic acid comprising: (a) an enzymatic cleavage domain comprising a single-stranded region, said single-stranded region comprising at least one internucleotide linkage 3' to an adenosine residue, at least one internucleotide linkage 3' to a cytosine residue, at least one internucleotide linkage 3' to a guanosine residue, and at least one internucleotide linkage 3' to a uridine residue, and wherein said enzymatic cleavage domain does not comprise a deoxyribonuclease-cleavable internucleotide linkage; (b) a fluorescence reporter group on one side of the internucleotide linkages; and (c) a non-fluorescent fluorescence-quenching group on the other side of the internucleotide linkages.
2. (Previously presented) The nucleic acid of claim 1, wherein the fluorescence-quenching group is a nitrogen-substituted xanthene compound, a substituted 4-(phenyldiazenyl)phenyl amine compound, or a substituted 4-(phenyldiazenyl)naphthylamine compound.
3. (Previously presented) The nucleic acid of claim 1, wherein the fluorescence-quenching group is 4-(4'-dimethylaminophenylazo)benzoic acid), N,N'-dimethyl-N,N'-diphenyl-4-((5-t-butoxycarbonylaminopentyl) aminocarbonyl) piperidinylsulfonerhodamine, 4',5'-dinitrofluorescein, piperic acid amide, 4-[4-nitrophenyldiazinyl]phenylamine, or 4-[4-nitrophenyldiazinyl]naphthylamine.
4. (Previously presented) The nucleic acid of claim 1, wherein the fluorescence reporter group is fluorescein, tetrachlorofluorescein, hexachlorofluorescein, rhodamine, tetramethylrhodamine, a Cy dye, Texas Red, a Bodipy dye, or an Alexa dye.
5. (Previously presented) The nucleic acid of claim 1, wherein the fluorescence reporter group is attached to the 5'-terminal nucleotide of the nucleic acid.

6. (Previously presented) The nucleic acid of claim 1, wherein the fluorescence quenching group is attached to the 5'-terminal nucleotide of the nucleic acid.
7. (Previously presented) The nucleic acid of claim 1 which is a single-stranded RNA molecule.
8. (Previously presented) The nucleic acid of claim 1 which is a chimeric oligonucleotide comprising a nuclease resistant modified ribonucleotide residue.
9. (Previously presented) The nucleic acid of claim 8, wherein the modified ribonucleotide residue is an 2'-O-methyl ribonucleotide, a 2'-methoxyethoxy ribonucleotide, a 2'-O-allyl ribonucleotide, a 2'-O-pentyl ribonucleotide, or a 2'-O-butyl ribonucleotide.
10. (Currently amended) The nucleic acid of claim 8, wherein the modified ribonucleotide residue is at the 5'-terminus or the 3'-terminus of the enzymatic cleavage domain.
11. (Previously presented) The nucleic acid of claim 1 which is a chimeric oligonucleotide comprising a deoxyribonuclease resistant modified deoxyribonucleotide residue.
12. (Previously presented) The nucleic acid of claim 11, wherein the deoxyribonuclease resistant modified deoxyribonucleotide residue is a phosphotriester deoxyribonucleotide, a methylphosphonate deoxyribonucleotide, a phosphoramidate deoxyribonucleotide, a phosphorothioate deoxyribonucleotide, a phosphorodithioate deoxyribonucleotide, or a 5 boranophosphate deoxyribonucleotide.
13. (Currently amended) The nucleic acid of claim 11, wherein the deoxyribonuclease resistant modified deoxyribonucleotide residue is in the enzymatic

cleavage domain.

14. (Previously presented) The nucleic acid of claim 1 which comprises a ribonuclease-cleavable modified ribonucleotide residue.

15. (Currently amended) The nucleic acid of claim 14, wherein the ribonuclease-cleavable modified ribonucleotide residue is in the enzymatic cleavage domain.

16. (Previously presented) The nucleic acid of claim 1 which is 5-30 nucleotides in length.

17. (Previously presented) The nucleic acid of claim 16 which is 7-10 nucleotides in length.

18. (Currently amended) The nucleic acid of claim 1, wherein the fluorescence-quenching group is 5' to the enzymatic cleavage domain and the fluorescence reporter group is 3' to the enzymatic cleavage domain.

19. (Previously presented) The nucleic acid of claim 18, wherein the fluorescence-quenching group is at the 5' terminus of the nucleic acid.

20. (Previously presented) The nucleic acid of claim 18, wherein the fluorescence reporter group is at the 3' terminus of the nucleic acid.

21. (Currently amended) The nucleic acid of claim 1, wherein the fluorescence reporter group is 5' to the enzymatic cleavage domain and the fluorescence-quenching group is 3' to the enzymatic cleavage domain.

22. (Previously presented) The nucleic acid of claim 21, wherein the fluorescence reporter group is at the 5' terminus of the nucleic acid.

23. (Previously presented) The nucleic acid of claim 21, wherein the fluorescence-quenching group is at the 3' terminus of the nucleic acid.

24. (Currently amended) The nucleic acid of claim 1, in which the enzymatic cleavage domain comprises the formula: 5'-N₁-n-N₂-3', wherein: (a) "N₁" represents zero to five 2'-modified ribonucleotide residues; (b) "N₂" represents one to five 2'-modified ribonucleotide residues; and (c) "n" represents four to ten unmodified ribonucleotide residues.

25. (Previously presented) The nucleic acid of claim 24, wherein the fluorescence-quenching group "N₁" represents one to five 2'-modified ribonucleotide residues.

26. (Previously presented) The nucleic acid of claim 25, wherein the fluorescence-quenching group is attached to the 5'-terminal 2'-modified ribonucleotide residue of N.sub.1.

27. (Previously presented) The nucleic acid of claim 25, wherein the fluorescence reporter group is attached to the 5'-terminal 2'-modified ribonucleotide residue of N₁.

28. (Currently amended) The nucleic acid of claim 24, wherein the fluorescence-quenching group is 5' to the enzymatic cleavage domain and the fluorescence reporter group is 3' to the enzymatic cleavage domain.

29. (Currently amended) The nucleic acid of claim 24, wherein the fluorescence reporter group is 5' to the enzymatic cleavage domain and the fluorescence-quenching group is 3' to the enzymatic cleavage domain.

30. (Currently amended) The nucleic acid of claim 24, wherein the enzymatic cleavage domain comprises the sequence "auggc".

31. (Previously presented) The nucleic acid of claim 30, wherein N₁ and N₂ each

represent one 2'-modified ribonucleotide residue.

32. (Previously presented) The nucleic acid of claim 31, wherein the 2'-modified ribonucleotide residue is an adenosine.

33. (Currently amended) A kit comprising: (a) in one container, a substrate, said substrate comprising a nucleic acid molecule comprising a single stranded region, said single-stranded region comprising i. an enzymatic cleavage domain comprising a single-stranded region, said single-stranded region comprising at least one internucleotide linkage 3' to an adenosine residue, at least one internucleotide linkage 3' to a cytosine residue, at least one internucleotide linkage 3' to a guanosine residue, and at least one internucleotide linkage 3' to a uridine residue, and wherein said enzymatic cleavage domain does not comprise a deoxyribonuclease cleavable internucleotide linkage; ii. a fluorescence reporter group on one side of the internucleotide linkages; and iii. a non-fluorescent fluorescence-quenching group on the other side of the internucleotide linkages.

34. (Previously presented) The kit of claim 33, further comprising in a second container a ribonuclease.

35. (Previously presented) The kit of claim 33, further comprising ribonuclease-free water.

36. (Previously presented) The kit of claim 33, further comprising a buffer.

37. (Previously presented) The kit of claim 33, further comprising ribonuclease-free laboratory plasticware.

38. (Previously presented) A method for measuring the activity of a ribonuclease comprising the steps of obtaining a sample from which the activity of the ribonuclease is to be measured, mixing the sample with the nucleic acid of claim 1, measuring the

amount of fluorescence that is produced, and correlating the amount of fluorescence that is produced to the activity of the ribonuclease.

39. (Previously presented) The method of claim 38 wherein the step for measuring the amount of fluorescence produced is carried out by measuring fluorescence in a fluorimeter.